AMENDMENTS TO THE CLAIMS

Listing of Claims:

(Cancelled) Claims 1-79

80. (New) A bidentate motif capable of binding a cytoplasmic protein and activating cellular

activities in a cell, said bidentate motif comprising a tyrosine and a serine/threonine residue

which are capable of interaction with cytoplasmic proteins, and wherein the residue and

cytoplasmic protein can interact to activate cellular activity in the cell.

81. (New) A bidentate motif according to claim 80 wherein the tyrosine and serine/threonine

residue comprises a binary switch for independent regulation of cellular activity.

82. (New) A bidentate motif capable of binding to a cytoplasmic protein according to claim 80

comprising a tyrosine and a serine/threonine residue, said motif comprising an amino acid

sequence alignment selected from the group consisting of:

$$N-X-X-Y-(X)_{1-13}-[R/K/H/Q]-[X/\psi]_{2-3}-S/T-X-P$$

wherein X is any residue, Y is tyrosine, SIT is serine or threonine and T is a hydrophibic residue

or an equivalent thereof; or

$$\underline{Y}$$
-(X) ₁₋₁₆-[R/K/H/Q]-[X/ ψ]₂₋₃- \underline{S} / \underline{T} -X-P

wherein X is any residue, Y is tyrosine, S/T is serine or threonine and T is a hydrophibic residue

or an equivalent thereof; or

$$N\text{-}X\text{-}X\text{-}\underline{Y}\text{-}[X]_{1\text{-}30}\text{-}[R/K/Q'H]\text{-}[X]_{1\text{-}4}\text{-}[\underline{S}/\underline{T}]\text{-}X\text{-}p$$

wherein X is any residue, Y is phosphotyrosine, $\underline{S}/\underline{T}$ is phosphoserine/phosphothreonins.

83. (New) A bidentate motif according to claim 80 wherein the motif is derived from a receptor.

84. (New) A bidentate motif according to claim 80 wherein the motif is derived from the

common beta chain (βc).

85. (New) A bidentate motif according to claim 80 wherein the tyrosine is equivalent to Tyr577 of the

common beta chain (β c) and/or the serine is equivalent to Ser 585 of the common beta chain (β c).

86. (New) A bidentate motif according to claim 80 wherein the tyrosine or serine/threonine are

independently phosphorylated in response to a cytokine, and phosphorylation is dependent on the

cytokine concentration.

87. (New) A bidentate motif according to claim 80 wherein phosphorylation of the serine

independently of the tyrosine regulates cell survival.

88. (New) A bidentate motif according to claim 80 wherein phosphorylation of the tyrosine

independent of the serine regulates cell survival and proliferation.

89. (New) A bidentate motif according to claim 83, with a modification at a residue equivalent to the

Tyr 577 and/or Ser585.

90. (New) The bidentate motif according to claim 89 wherein the residue equivalent to Tyr 577 is

substituted with phenylalanine and/or the Ser 585 residue is substituted with glycine.

91. (New) A method of modulating cellular activity in a cell, said method comprising: modulating

phosphorylation of a tyrosine and/or serine residue of a bidentate motif capable of binding to a

cytoplasmic protein comprising a tyrosine and a serinelthreonine residue, said motif comprising an

amino acid sequence alignment selected from the group consisting of:

 $N-X-X-Y-(X)_{1-13}-[R/K/H/Q]-[X/\psi]_{2-3}-S/T-X-P$

wherein X is any residue, Y is tyrosine, SIT is serine or threonine and 'F is a hydrophibic residue or an equivalent thereof; or

$$\underline{Y}$$
- $(X)_{1-16}$ - $[R/K/H/Q)$ - $[X/\psi]_{2-3}$ - $\underline{S}/\underline{T}$ - X - P

wherein X is any residue, Y is tyrosine, S/T is serine or threonine and T is a hydrophibic residue or an equivalent thereof; or

$$N-X-X-Y-[X]_{1-30}-[R/K/Q/H]-[X]_{1-4}[S/T]-X-p$$

wherein X is any residue, Y is phosphotyrosine, S/T is phosphoserine/phosphothreonine.

- 92. (New) A method according to claim 91 wherein the phosphorylation is modulated by mutating the tyrosine and/or serine.
- 93. (New) A method according to claim 92 wherein the Tyr is substituted for phenylalanine and/or the serine is substituted for glycine.
- 94. (New) A method according to claim 91 wherein the phosphorylation is decreased by subjecting the cell to an antagonist or kinase inhibitor which inhibits phosphorylation of the tyrosine and/or serine.
- 95. (New) A method according to claim 91 wherein cellular activity is inhibited, said method comprising decreasing or inhibiting phosphorylation of the tyrosine andlor serine of the bidentate motif.
- 96. (New) A method according to claim 95 wherein the cellular activity is cell survival, said method comprising inhibiting phosphorylation of the serine.
- 97. (New) A method according to claim 95 wherein the cellular activity is cell survival, said method comprising inhibiting phosphorylation of the serine equivalent to Ser585 of the common βc .
- 98. (New) A method according to claim 91 wherein cellular activity is activated, said method comprising inducing phosphorylation of the tyrosine and/or serine of the bidentate motif.

Application Serial No.: 10/595,562 Docket No.: 03991/0204242-US0 Page 4

99. (New) A method according to claim 98 wherein the cellular activity is cell survival, said method comprising increasing phosphorylation of the serine.

100. (New) A method according to claim 91 wherein the cellular activity is cell proliferation, said method comprising increasing phosphorylation of the tyrosine.

101. (New) A method of treating a cytokine mediated condition, said method comprising:

regulating activation of phosphorylation of a tyrosine and/or serine of a bidentate motif capable of binding to a cytoplasmic protein comprising a tyrosine and a serine/threonine residue, said motif comprising an amino acid sequence alignment selected from the group consisting of:

$$N-X-X-Y-(X)_{1-13}-[R/K/H/Q]-[X/\psi]_{2-3}-S/T-X-P$$

wherein X is any residue, Y is tyrosine, SIT is serine or threonine and T is a hydrophibic residue or an equivalent thereof; or

$$\underline{Y}$$
- $(X)_{1-16}$ - $[R/K/H/Q]$ - $[X/\psi]_{2-3}$ - $[\underline{S/T}]$ - X - P

wherein X is any residue, Y is tyrosine, S/T is serine or threonine and ψ is a hydrophibic residue or an equivalent thereof; or

$$N - X - X - \underline{Y} - [X]_{1 - 30} - [R/K/Q/H] - [X]_{1 - 4} - [\underline{S/T}] - X - p$$

wherein X is any residue, Y is phosphotyrosine, S/T is phosphoserine/phosphothreonine.

102. (New) A method according to claim 101 wherein the cytokine mediated condition is treated by increasing or decreasing activation of phosphorylation of the tyrosine and/or serine of the bidentate motif.

103. (New) A method according to claim 101 wherein the phosphorylation is decreased by mutating the tyrosine and/or serine.

104. (New) A method use according to claim 103 wherein the motif is mutated by substituting tyrosine

for phenylalanine and/or substituting serine for glycine.

105. (New) A method according to claim 101 wherein the phosphorylation is decreased by subjecting

the cell to an antagonist which inhibits phosphorylation of the tyrosine and/or serine.

106. (New) A method according to claim 101 wherein the cytokine mediated condition is a GM-CSF

mediated condition.

107. (New) A method according to claim 101 wherein the cytokine mediated condition involves cell

survival.

108. (New) A method according to claim 101 wherein the cytokine mediated condition involves cell

proliferation.

109. (New) A method according to claim 101 wherein the cytokine mediated condition is selected

from the group consisting of myeloid cell activation, asthma and rheumatoid arthritis.

110. (New) A method for diagnosing a proliferative condition involving cell proliferation or cell

survival, said method including:

detecting a level of phosphorylation of tyrosine and/or serine in a bidentate motif capable of

binding to a cytoplasmic protein comprising a tyrosine and a serine/threonine residue, said motif

comprising an amino acid sequence alignment selected from the group consisting of:

$$N-X-X-Y-(X)_{1-13}-[R/K/H/Q]-[X/\psi]_{2-3}-S/T-X-P$$

wherein X is any residue, Y is tyrosine, SIT is serine or threonine and T is a hydrophibic residue or an

equivalent thereof; or

 \underline{Y} -(X)₁₋₁₆-[R/K/H/Q]-[X/ ψ]₂₋₃- \underline{S} / \underline{T} -X-P

wherein X is any residue, Y is tyrosine, S/T is serine or threonine and ψ is a hydrophibic residue or an equivalent thereof; or

$$N\hbox{-} X\hbox{-} X\hbox{-} \underline{Y}\hbox{-} [X]_{1\hbox{-}30}\hbox{-} [R/K/Q/H]\hbox{-} [X]_{1\hbox{-}4}\hbox{-} [\underline{S}/\underline{T}]\hbox{-} X\hbox{-} p$$

wherein X is any residue, Y is phosphotyrosine, S/T is phosphoserine/phosphothreonine; and comparing against a cell of a normal level of phosphorylation.

Application Serial No.: 10/595,562 Docket No.: 03991/0204242-US0